WO 98/40747 Attorney Name. Katherine M. Kowalchyk (Reg. No. 36,848) Phone No. 612 371 5311

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FIGURE 1

```
PRI
        LOCUS
                                            HSTGFB3M
                                                                          2574 bp
                                                                                                   RNA
                                                                                                                                                         12-SEP-1993
                                            Human mRNA for transforming growth factor-beta 3 (TGF-beta
        DEFINITION
                                             3).
                                             X14149
        ACCESSION
                                             g37095
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                                             growth factor; transforming growth factor; transforming
                                             growth factor-beta 3.
        SOURCE
                                             human.
Agency Control of the Control of the
             ORGANISM
                                             Homo sapiens
                                             Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;
                                             Vertebrata: Eutheria: Primates: Catarrhini: Hominidae;
 ū
Ŀ.
        REFERENCE
                                             1 (bases 1 to 2574)
L
             AUTHORS
                                             Chen, E.Y.
ũ
             TITLE
                                             Direct Submission
                                             Submitted (23-MAR-1989) Chen E.Y., Genentech Inc., 460 Pt.
             JOURNAL
=
£.
                                             San Bruno Blvd., San Francisco, CA 94080, USA
N
        REFERENCE
                                                    (bases 1 to 2574)
AUTHORS
                                             Derynck, R., Lindquist, P.B., Lee, A., Wen, D., Tamm, J.,
                                             Graycar, J.L., Rhee, L., Mason, A.J., Miller, D.A.,
                                             Coffey, R.J., Moses, H.L. and Chen, E.Y.
             TITLE
                                             A new type of transforming growth factor-beta, TGF-beta 3
                                             EMBO J. 7 (12), 3737-3743 (1988)
             JOURNAL
             MEDLINE
                                             89091120
                                             See <J03241> for alternative sequence of TGF-beta 3.
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          BASE COUNT
                                               629 a
                                                                   680 c
                                                                                       666 g
                                                                                                            599 t
```

ORIGIN

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FIGURE 1 (cont'd)

```
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 121 cacettettg ccaageetca gtetttggga tetggggagg ccgeetggtt tteeteete
 181 cttctgcacg tctgctgggg tctcttcctc tccaggcctt gccgtccccc tggcctctct
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 901 gtccaactta ggtctagaaa tcagcattca ctgtccatgt cacacctttc agcccaatgg
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1381 ttgctgcgtg ccccaggacc tggagcccct gaccatcctg tactatgttg ggaggacccc
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1801 tggcaatggg atttgggcta ccctaaaggg agaaggaagg gcagagaatg gctgggtcag
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2161 gaaagggtgg aaatcaaccc tctcctgtct gccctctggg tccctctct cacctctccc
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2281 tggattgtgg ttccatgcag ccttggggca ttatgggtct tcccccactt cccttcaag
2341 accordate cattleggest tootegeage aggregatea acategeage cattleggesa
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2521 atttcatctt ctggaatatt gtttctagaa cagtaaaagc cttattctaa ggtg
11
```

Inventor: CANIGGIA et al. Docket No: 11757.38USD1 Title: METHODS TO DIAGNOSE A REQUIRED REGULATION OF TROPHOBLAST INVASION

WO 98/40747 Phone No. 612.371 5311

Attorney Name. Katherine M Kowalchyk (Reg No. 36,848)

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FIGURE 2

PRI

```
28-JUN-1995
                            3678 bp
                                       mRNA
               HSU22431
  LOCUS
              Human hypoxia-inducible factor 1 alpha (HIF-1 alpha) mRNA, complete
  DEFINITION
               cds.
               U22431
  ACCESSION
               g881345
  NID
KEYWORDS
  SOURCE
               human.
              Homo sapiens
    ORGANISM
Eukaryotae; mitochondrial eukaryotes; Metazoa; Chordata;
Vertebrata; Eutheria; Primates; Catarrhini; Hominidae; Homo.
I
               1 (bases 1 to 3678)
  REFERENCE
               Wang, G.L., Jiang, B.H., Rue, E.A. and Semenza, G.L.
    AUTHORS
<u>_</u>
               Hypoxia-inducible factor 1 is a basic-helix-loop-helix-PAS
u
    TITLE
               heterodimer regulated by cellular O2 tension
IJ
               Proc. Natl. Acad. Sci. U.S.A. 92 (12), 5510-5514 (1995)
    JOURNAL
33
               95296340
    MEDLINE
1
                  (bases 1 to 3678)
   REFERENCE
               Wang, G.L., Jiang, B.-H., Rue, E.A. and Semenza, G.L.
T.
    AUTHORS
               Direct Submission
    TITLE
T.
               Submitted (09-MAR-1995) Gregg L. Semenza, Center for Medical
     JOURNAL
Genetics, The Johns Hopkins University School of Medicine, 600 N.
               Wolfe St., Baltimore, MD 21287-3914, USA
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                         CLVLICEPIPHPSNIEIPLDSKTFLSRHSLDMKFSYCDERITELMGYEPEELLGRSIY
                         EYYHALDSDHLTKTHHDMFTKGQVTTGQYRMLAKRGGYVWVETQATVIYNTKNSQPQC
                         IVCVNYVVSGIIQHDLIFSLQQTECVLKPVESSDMKMTQLFTKVESEDTSSLFDKLKK
                         EPDALTLLAPAAGDTIISLDFGSNDTETDDQQLEEVPLYNDVMLPSPNEKLQNINLAM
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                                              1111 t
                   1197 a
                              695 c
    BASE COUNT
```

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FIGURE 2 (cont'd)

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1 gtgaagacat cgcggggacc gattcaccat ggagggcgcc ggcggcgcga acgacaagaa
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 241 acttctggat gctggtgatt tggatattga agatgacatg aaagcacaga tgaattgctt
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 421 tgtgtttgat tttactcatc catgtgacca tgaggaaatg agagaaatgc ttacacacag
 481 aaatggcctt gtgaaaaagg gtaaagaaca aaacacacag cgaagctttt ttctcagaat
 541 gaagtgtacc ctaactagcc gaggaagaac tatgaacata aagtctgcaa catggaaggt
 601 attgcactgc acaggocaca ttcacgtata tgataccaac agtaaccaac ctcagtgtgg
 661 gtataagaaa ccacctatga cctgcttggt gctgatttgt gaacccattc ctcacccatc
 721 aaatattgaa attootttag atagcaagac tttootcagt cgacacagcc tggatatgaa
 781 attttcttat tgtgatgaaa gaattaccga attgatggga tatgagccag aagaactttt
 841 aggocgotca atttatgaat attatoatgo tttggactot gatoatotga coasaactoa
 901 tratgatatg titartaaag garaagtrar caraggarag taraggatge tigraaaag
 961 aggtggatat gtctgggttg aaactcaagc aactgtcata tataacacca agaattctca
1021 accacagige attgtatgig tgaattacgt tgtgagtggt attaticage acgacitgat
1081 tttctccctt caacaaacag aatgtgtcct taaaccggtt gaatcttcag atatgaaaat
1141 gactcageta ttcaccaaag ttgaatcaga agatacaagt agcetetttg acaaacttaa
1201 gaaggaacet gatgetttaa etttgetgge eccageeget ggagacacaa teatatettt
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3061 tggcctatga aattgttaaa cctggaacat gacattgtta atcatataat aatgattctt
3121 aaatgotgta tggtttatta tttaaatggg taaagccatt tacataatat agaaagatat
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3661 ataaacatct tctgtgga
```

Inventor: CANIGGIA et al.

Docket No 11757 38USD1

Title: METHODS TO DIAGNOSE A REQUIRED REGULATION OF TROPHOBLAST

INVASION

Attorney Name. Katherine M Kowalchyk (Reg. No 36,848)

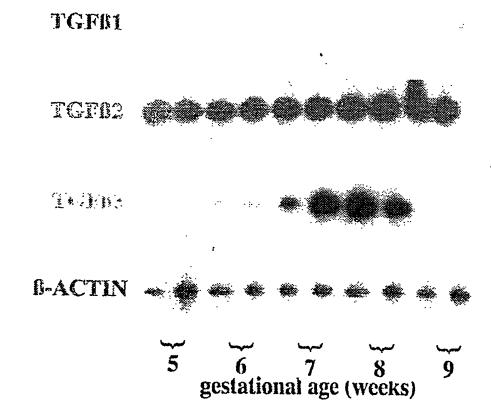
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FIGURE 3A



Inventor: CANIGGIA et al.
Docket No · 11757 38USD1
Title METHODS TO DIAGNOSE A REQUIRED REGULATION

Attorney Name. Katherine M Kowalchyk (Reg. No. 36,848) Phone No.: 612.371 5311

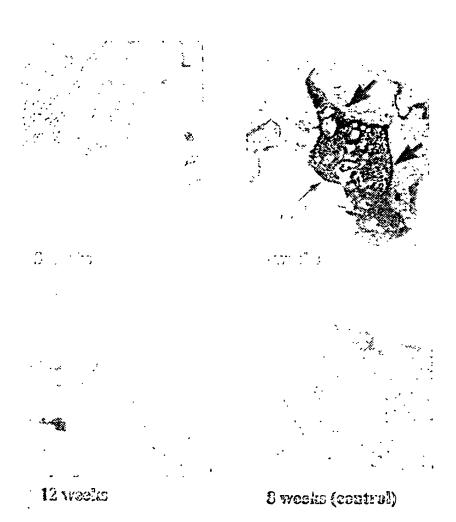
and the first than the after much than

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FIGURE 3B

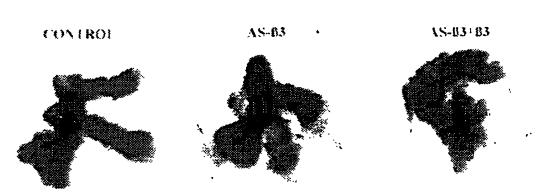


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FIGURE 4A



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Docket No 11757.38USD1
Title: METHODS TO DIAGNOSE A REQUIRED REGULATION OF TROPHOBLAST INVASION Attorney Name Katherine M. Kowalchyk (Reg No 36,848) Phone No · 612 371.5311

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FIGURE 4B

S-B3

AS-83

AS-B3+B3



200

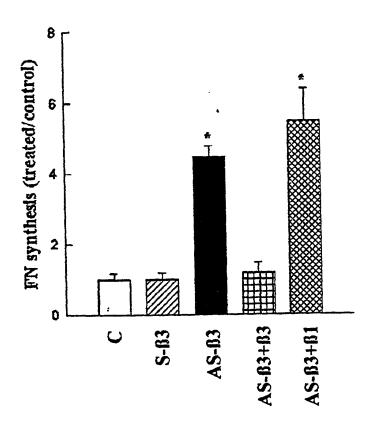
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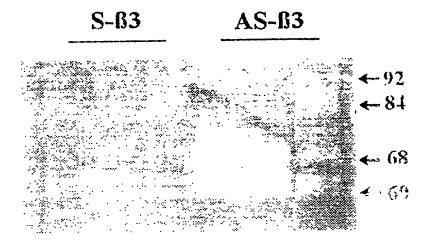
FIGURE 4C



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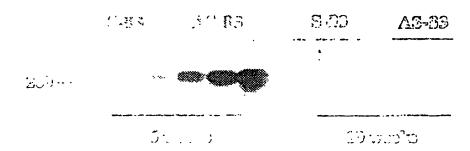
FIGURE 4D



Inventor: CANIGGIA et al. Docket No.: 11757 38USD1 Title: METHODS TO DIAGNOSE A REQUIRED REGULATION OF TROPHOBLAST INVASION Attorney Name. Katherine M Kowalchyk (Reg. No. 36,848)
Phone No.: 612 371 5311
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Title METHODS TO DIAGNOSE A REQUIRED REGULATION
OF TROPHOBLAST INVASION
Attorney Name Katherine M Kowalchyk (Reg No 36,848)
Phone No · 612.371 5311
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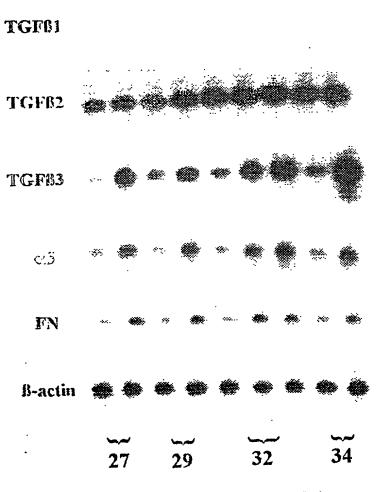
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FIGURE 5A

PE C PE PE C PE C PE



gestational age (weeks)

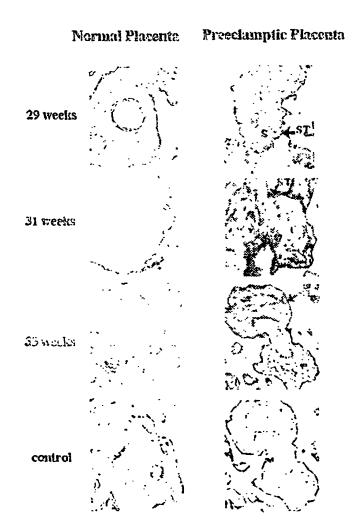
Attorney Name: Katherine M Kowalchyk (Reg No. 36,848)

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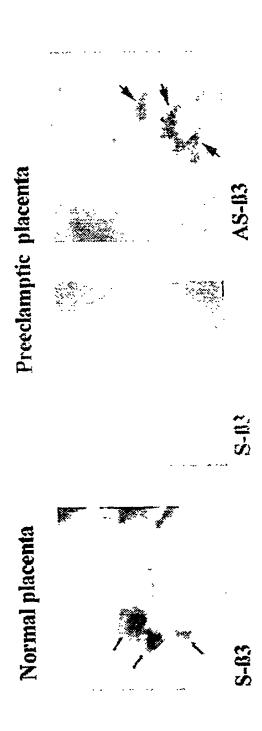
FIGURE 5B



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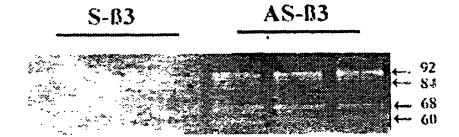




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FIGURE 6B



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Docket No · 11757.38USD1
Title METHODS TO DIAGNOSE A REQUIRED REGULATION
OF TROPHOBLAST INVASION Attorney Name: Katherine M. Kowalchyk (Reg. No. 36,848)
Phone No. 612 371 5311

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FIGURE 6C

S-B3

AS-B3

Inventor: CANIGGIA et al. Docket No.: 11757.38USD1 Title METHODS TO DIAGNOSE A REQUIRED REGULATION OF TROPHOBLAST INVASION

Attorney Name: Katherine M Kowalchyk (Reg No 36,848) Phone No. 612 371.5311

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FIGURE 7A

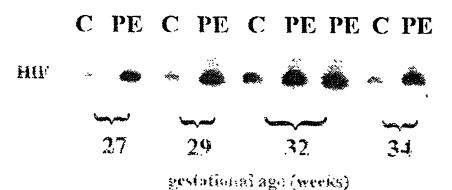
locets ettos

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FIGURE 7B



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FIGURE 8

20% 3%

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B-ACTIN

SUBSTITUTE SHEET (RULE 26)

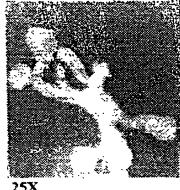
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20/21

FIGURE 9

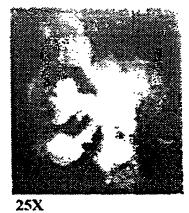
20% O2

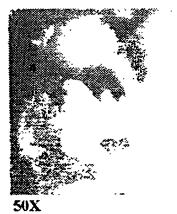




50X

3% O2





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FIGURE 10

